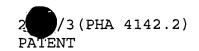


# REMARKS

The purpose of the instant amendment is to provide chemical names for the trademarks or trade names contained in the application, thereby conforming this application to the parent application, Serial No. 08/661,641. No new matter has been added by this amendment. Favorable consideration and early allowance of all pending claims is requested.



# VERSION WITH MARKINGS TO SHOW CHANGES MADE

# IN THE RELATED CASE:

Please replace the paragraph on Page 1, lines 7-8 with the following:

--This is a <u>continuation of US Application Serial No. 08/661,641,</u>
<u>filed June 11, 1996 which application is a</u> continuation-in-part
of US Application Serial No. 08/489,415, filed June 12, 1995.--

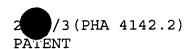
# IN THE SPECIFICATION:

Please replace the paragraph on page 5, lines 6-12 with the following:

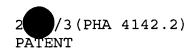
--In addition, the invention describes a combination comprising a therapeutically-effective amount of a leukotriene B4 receptor antagonist and a cyclooxygenase-2 inhibitor selected from Taisho NS-398 (Methanesulfonamide, N-[2-(cyclohexyloxy)-4-nitrophenyl]), meloxicam (2H-1,2-Benzothiazine-3-carboxamide, 4-hydroxy-2-methyl-N-(5-methyl-2-thiazolyl)-, 1,1-dioxide), [floculide] flosulide (Methanesulfonamide, N-[6-(2,4-difluorophenoxy)-2,3-dihydro-1-oxo-1H-inden-5-yl]), Merck MK-966 (2(5H)-Furanone, 4-[4-(methylsulfonyl)phenyl]-3-phenyl), Merck L-752,860 and compounds of Formula I--

Please replace the paragraph beginning on page 8, lines 26-37 and ending on page 9, lines 1-5 with the following:

--Preferred leukotriene B<sub>4</sub> receptor antagonists include Bayer
Bay-x-1005 ((R)-a-Cyclopentyl-4-(2-quinolinylmethoxy) benzene
acetic acid), Ciba-Geigy CGS-25019C (Benzamide, 4-[[5-[4(aminoiminomethyl)phenoxy]pentyl]oxy]-3-methoxy-N,N-bis(1methylethyl)-, (2Z)-2-butenedioate), ebselen (1,2Benzisoselenazol-3(2H)-one, 2-phenyl), Leo Denmark ETH-615
(Benzoic acid, 4-[[[(3-fluorophenyl)methyl][4-(2quinolinylmethoxy)phenyl]amino]methyl]), Lilly LY-293111 (Benzoic



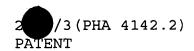
acid, 2-[3-[3-[(5-ethyl-4'-fluoro-2-hydroxy[1,1'-biphenyl]-4yl)oxy[propoxy]-2-propylphenoxy]), Ono ONO-4057 (Benzenepropanoic acid, 2-(4-carboxybutoxy)-6-[[(5E)-6-(4-methoxyphenyl)-5hexenyl]oxy]), Terumo TMK-688 (Carbonic acid, 4-[5-[[2-[4-(diphenylmethoxy) -1-piperidinyl]ethyl]amino]-5-oxo-1,3pentadienyl]-2-methoxyphenyl ethyl ester), Boehringer Ingleheim BI-RM-270 (2-Benzoxazolamine, N-[(1S)-2-cyclohexyl-1-(2pyridinyl)ethyl]-5-methyl), Lilly LY 213024 (Benzenepropanoic acid, 5-(3-carboxybenzoyl)-2-(decyloxy)), Lilly LY 264086 (9H-Xanthene-4-propanoic acid, 7-carboxy-3-(decyloxy)-9-oxo), Lilly LY 292728, Ono ONO LB457 (Benzenepropanoic acid, 2-(4carboxybutoxy) -6-[[(5E) -6-(4-methoxyphenyl) -5-hexenyl]oxy], Pfizer 105696, Perdue Frederick PF 10042 (Pyrrolidine, 1-[5hydroxy-5-[8-(1-hydroxy-2-phenylethyl)-2-dibenzofuranyl]-1oxopentyl]), Rhone-Poulenc Rorer RP 66153 (2-Thiopheneheptanoic acid, .alpha.,.alpha.-dimethyl-3-(3-phenylpropyl)), SmithKline Beecham SB-201146 (2-Propenoic acid, 3-[6-[[(3aminophenyl)sulfinyl]methyl]-3-[[8-(4-methoxyphenyl)octyl]oxy]-2-pyridinyl]-, (2E)), SmithKline Beecham SB-201993 (Benzoic acid, 3-[[[[6-[(1E)-2-carboxyethenyl]-5-[[8-(4methoxyphenyl)octyl]oxy]-2-pyridinyl]methyl]thio]methyl]), Searle SC-53228 (2H-1-Benzopyran-2-propanoic acid, 7-[3-[2-(cyclopropylmethyl) - 3 - methoxy - 4 -[(methylamino)carbonyl]phenoxy]propoxy]-3,4-dihydro-8-propyl-, (2S)), Sumitamo SM 15178 (.beta.-Alanine, N-[[6-[(4-acetyl-2ethyl-5-hydroxyphenoxy)methyl]-2- pyridinyl]carbonyl]-N-ethyl), American Home Products WAY 121006 ([1,1'-Biphenyl]-4-acetic acid, 2-fluoro-4'-(2-quinolinylmethoxy)), Bayer Bay-o-8276, calcitriol (9,10-Secocholesta-5,7,10(19)-triene-1,3,25-triol, (1.alpha., 3.beta., 5Z, 7E)), Warner-Lambert CI-987 (2,4-Thiazolidinedione, 5-[[3,5-bis(1,1-dimethylethyl)-4hydroxyphenyl]methylene]), Merck and Co. L-651392 (3H-Phenothiazin-3-one, 4-bromo-2,7-dimethoxy), Lilly LY 210073, Lilly LY 223982 (Benzenepropanoic acid, 5-(3-carboxybenzoyl)-2-[[(5E)-6-(4-methoxyphenyl)-5- hexenyl]oxy]), Lilly LY-233569 (2-Propenamide, N-hydroxy-N-methyl-3-[2-(methylthio)phenyl]), Lilly



LY-255283 (Ethanone, 1-[5-ethyl-2-hydroxy-4-[[6-methyl-6-(1Htetrazol-5-yl)heptyl]oxy]phenyl]), Merck and Co. MK-591 (1H-Indole-2-propanoic acid, 1-[(4-chlorophenyl)methyl]-3-[(1,1dimethylethyl)thio]-a,a-dimethyl-5-(2-quinolinylmethoxy)-, sodium salt), Merck and CO. MK-886 (1H-indole-2-propanoic acid, 1-[(4chlorophenyl)methyl]-3-[(1,1-dimethylethyl)thio]-a,a-dimethyl-5-(1-methylethyl)), Ono ONO-LB-448, Purdue Frederick PF-5901 (Benzenemethanol, a-pentyl-3-(2-quinolinylmethoxy)), Rhone-Poulenc Rorer RG 14893 (2-Naphthalenecarboxylic acid, 4-[2-[methyl(2-phenylethyl)amino]-2-oxoethyl]-8-(phenylmethoxy)), Rhone-Poulenc Rorer RP 66364, Rhone-Poulenc Rorer RP 69698 (Pyridine, 2-[[5-methyl-5-(1H-tetrazol-5-yl)hexyl]oxy]-4,6diphenyl), Searle SC-41930 (2H-1-Benzopyran-2-carboxylic acid, 7-[3-(4-acetyl-3-methoxy-2-propylphenoxy)propoxy]-3,4-dihydro-8propyl), Searle SC-50505, Searle SC-51146, SmithKline Beecham SK&F-104493 (5H-Pyrrolo[1,2-a]imidazole, 6,7-dihydro-2-(4methoxyphenyl) - 3 - (4 - pyridinyl)), and Teinjin TEI - 1338 (Benzoic acid, 2-[[4-[2-[2-(2-naphthalenyl)ethenyl]cyclopropyl]-1oxobutyl]amino]-, methyl ester, [1R-[1.alpha.,2.beta.(E)]]).--

Please replace the paragraph on page 9, lines 6-15 with the following:

--More preferred leukotriene B4 receptor antagonists include
Bayer Bay-x-1005 ((R)-a-Cyclopentyl-4-(2quinolinylmethoxy)benzeneacetic acid), Ciba-Geigy CGS-25019C
(Benzamide, 4-[[5-[4-(aminoiminomethyl)phenoxy]pentyl]oxy]-3methoxy-N,N-bis(1-methylethyl)-, (2Z)-2-butenedicate), ebselen
(1,2-Benzisoselenazol-3(2H)-one, 2-phenyl), Leo Denmark ETH-615
(Benzoic acid, 4-[[[(3-fluorophenyl)methyl][4-(2quinolinylmethoxy)phenyl]amino]methyl]), Lilly LY-293111 (Benzoic acid, 2-[3-[3-[(5-ethyl-4'-fluoro-2-hydroxy[1,1'-biphenyl]-4yl)oxy]propoxyl-2-propylphenoxyl), Ono ONO-4057 (Benzenepropanoic acid, 2-(4-carboxybutoxy)-6-[[(5E)-6-(4-methoxyphenyl)-5hexenyl]oxyl), Terumo TMK-688 (Carbonic acid, 4-[5-[[2-[4(diphenylmethoxy)-1-piperidinyl]ethyl]amino]-5-oxo-1,3pentadienyl]-2-methoxyphenyl ethyl ester), Boehringer Ingleheim

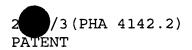


BI-RM-270 (2-Benzoxazolamine, N-[(1S)-2-cyclohexyl-1-(2pyridinyl)ethyl]-5-methyl), Lilly LY 213024 (Benzenepropanoic acid, 5-(3-carboxybenzoyl)-2-(decyloxy)), Lilly LY 264086 (9H-Xanthene-4-propanoic acid, 7-carboxy-3-(decyloxy)-9-oxo), Lilly LY 292728, Ono ONO LB457 (Benzenepropanoic acid, 2-(4carboxybutoxy) -6-[[(5E) -6-(4-methoxyphenyl) -5-hexenyl]oxy]), Pfizer 105696, Perdue Frederick PF 10042 (Pyrrolidine, 1-[5hydroxy-5-[8-(1-hydroxy-2-phenylethyl)-2-dibenzofuranyl]-1oxopentyl]), Rhone-Poulenc Rorer RP 66153 (2-Thiopheneheptanoic acid, .alpha.,.alpha.-dimethyl-3-(3-phenylpropyl)), SmithKline Beecham SB-201146 (2-Propenoic acid, 3-[6-[[(3aminophenyl)sulfinyl]methyl]-3-[[8-(4-methoxyphenyl)octyl]oxy]-2-pyridinyl]-, (2E)), SmithKline Beecham SB-201993 (Benzoic acid, methoxyphenyl)octyl]oxy]-2-pyridinyl]methyl]thio]methyl]), Searle SC-53228 (2H-1-Benzopyran-2-propanoic acid, 7-[3-[2-(cyclopropylmethyl) - 3-methoxy-4-[(methylamino)carbonyl]phenoxy]propoxy]-3,4-dihydro-8-propyl-, (2S)), Sumitamo SM 15178 (.beta.-Alanine, N-[[6-[(4-acetyl-2ethyl-5-hydroxyphenoxy)methyl]-2- pyridinyl]carbonyl]-N-ethyl), and American Home Products WAY 121006 ([1,1'-Biphenyl]-4-acetic

Please replace the paragraph on page 9, lines 16-19 with the following:

acid, 2-fluoro-4'-(2-quinolinylmethoxy)).--

--Even more preferred leukotriene B<sub>4</sub> receptor antagonists include Bayer Bay-x-1005 ((R)-a-Cyclopentyl-4-(2-quinolinylmethoxy)benzeneacetic acid), Ciba-Geigy CGS-25019C (Benzamide, 4-[[5-[4-(aminoiminomethyl)phenoxy]pentyl]oxy]-3-methoxy-N,N-bis(1-methylethyl)-, (2Z)-2-butenedioate), ebselen (1,2-Benzisoselenazol-3(2H)-one, 2-phenyl), Leo Denmark ETH-615 (Benzoic acid, 4-[[[(3-fluorophenyl)methyl][4-(2-quinolinylmethoxy)phenyl]amino]methyl]), Lilly LY-293111 (Benzoic acid, 2-[3-[3-[(5-ethyl-4'-fluoro-2-hydroxy[1,1'-biphenyl]-4-yl)oxy]propoxy]-2-propylphenoxy]), Ono ONO-4057 (Benzenepropanoic acid, 2-(4-carboxybutoxy)-6-[[(5E)-6-(4-methoxyphenyl)-5-



hexenyl]oxy]), and Terumo TMK-688 (Carbonic acid, 4-[5-[[2-[4-(diphenylmethoxy)-1-piperidinyl]ethyl]amino]-5-oxo-1,3-pentadienyl]-2-methoxyphenyl ethyl ester).--

## IN THE CLAIMS:

#### CLAIM 2:

2. (once amended) A combination comprising a therapeutically-effective amount of a leukotriene B<sub>4</sub> receptor antagonist and a cyclooxygenase-2 inhibitor selected from Taisho NS-398 (Methanesulfonamide, N-[2-(cyclohexyloxy)-4-nitrophenyl]), meloxicam (2H-1,2-Benzothiazine-3-carboxamide, 4-hydroxy-2-methyl-N-(5-methyl-2-thiazolyl)-, 1,1-dioxide), [floculide] flosulide (Methanesulfonamide, N-[6-(2,4-difluorophenoxy)-2,3-dihydro-1-oxo-1H-inden-5-yl]), Merck MK-966 (2(5H)-Furanone, 4-[4-(methylsulfonyl)phenyl]-3-phenyl), Merck L-752,860 and compounds of Formula I

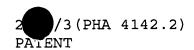
$$R^2$$
  $R^1$   $R^3$ 

wherein A is a substituent selected from partially unsaturated or unsaturated heterocyclyl and partially unsaturated or unsaturated carbocyclic rings;

wherein R<sup>1</sup> is at least one substituent selected from heterocyclyl, cycloalkyl, cycloalkenyl and aryl, wherein R<sup>1</sup> is optionally substituted at a substitutable position with one or more radicals selected from alkyl, haloalkyl, cyano, carboxyl, alkoxycarbonyl, hydroxyl, hydroxyalkyl, haloalkoxy, amino, alkylamino, arylamino, nitro, alkoxyalkyl, alkylsufinyl, halo, alkoxy and alkylthio;

wherein  ${\ensuremath{\mbox{R}}}^2$  is methyl or amino; and

wherein  $R^3$  is a radical selected from hydrido, halo, alkyl, alkenyl, alkynyl, oxo, cyano, carboxyl, cyanoalkyl,

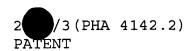


heterocyclyloxy, alkyloxy, alkylthio, alkylcarbonyl, cycloalkenyl, aralkyl, heterocyclylalkyl, acyl, alkylthioalkyl, hydroxyalkyl, alkoxycarbonyl, arylcarbonyl, aralkylcarbonyl, aralkenyl, alkoxyalkyl, arylthioalkyl, aryloxyalkyl, aralkylthioalkyl, aralkoxyalkyl, alkoxyaralkoxyalkyl, alkoxycarbonylalkyl, aminocarbonyl, aminocarbonylalkyl, alkylaminocarbonyl, N-arylaminocarbonyl, N-alkyl-N-arylaminocarbonyl, alkylamino, N-alkyl-N-aralkylamino, N-arylamino, N-arylamino, N-aralkylamino, N-alkyl-N-aralkylamino, N-alkyl-N-aralkylamino, N-alkyl-N-aralkylaminoalkyl, N-alkyl-N-aralkylaminoalkyl, N-alkyl-N-aralkylaminoalkyl, aryloxy, aralkoxy, arylthio, aralkylaminosulfonyl, alkylaminosulfonyl, aninosulfonyl, N-arylaminosulfonyl, arylsulfonyl, N-alkyl-N-arylaminosulfonyl, N-arylaminosulfonyl, arylsulfonyl, N-alkyl-N-arylaminosulfonyl;

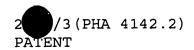
or a pharmaceutically-acceptable salt thereof.

### CLAIM 3:

3. (once amended) The combination of Claim 2 wherein the leukotriene B4 receptor antagonist is selected from Bayer Bay-x-1005 ((R)-a-Cyclopentyl-4-(2-quinolinylmethoxy) benzene acetic acid), Ciba-Geigy CGS-25019C (Benzamide, 4-[[5-[4-(aminoiminomethyl) phenoxy] pentyl] oxy] -3-methoxy-N, N-bis(1methylethyl)-, (2Z)-2-butenedioate), ebselen (1,2-Benzisoselenazol-3(2H)-one, 2-phenyl), Leo Denmark ETH-615 (Benzoic acid, 4-[[[(3-fluorophenyl)methyl][4-(2quinolinylmethoxy)phenyl]amino]methyl]), Lilly LY-293111 (Benzoic acid, 2-[3-[3-[(5-ethyl-4'-fluoro-2-hydroxy[1,1'-biphenyl]-4yl)oxy[propoxy]-2-propylphenoxy]), Ono ONO-4057 (Benzenepropanoic acid, 2-(4-carboxybutoxy)-6-[[(5E)-6-(4-methoxyphenyl)-5hexenyl]oxy]), Terumo TMK-688 (Carbonic acid, 4-[5-[[2-[4-(diphenylmethoxy) -1-piperidinyl]ethyl]amino]-5-oxo-1,3pentadienyl]-2-methoxyphenyl ethyl ester), Boehringer Ingleheim BI-RM-270 (2-Benzoxazolamine, N-[(1S)-2-cyclohexyl-1-(2pyridinyl)ethyl]-5-methyl), Lilly LY 213024 (Benzenepropanoic acid, 5-(3-carboxybenzoyl)-2-(decyloxy)), Lilly LY 264086 (9H-



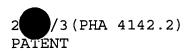
Xanthene-4-propanoic acid, 7-carboxy-3-(decyloxy)-9-oxo), Lilly LY 292728, Ono ONO LB457 (Benzenepropanoic acid, 2-(4carboxybutoxy) -6-[[(5E) -6-(4-methoxyphenyl) -5-hexenyl]oxy]), Pfizer 105696, Perdue Frederick PF 10042 (Pyrrolidine, 1-[5hydroxy-5-[8-(1-hydroxy-2-phenylethyl)-2-dibenzofuranyl]-1oxopentyl]), Rhone-Poulenc Rorer RP 66153 (2-Thiopheneheptanoic acid, .alpha.,.alpha.-dimethyl-3-(3-phenylpropyl)), SmithKline Beecham SB-201146 (2-Propenoic acid, 3-[6-[[(3aminophenyl)sulfinyl]methyl]-3-[[8-(4-methoxyphenyl)octyl]oxy]-2-pyridinyl]-, (2E)), SmithKline Beecham SB-201993 (Benzoic acid, 3-[[[6-[(1E)-2-carboxyethenyl]-5-[[8-(4methoxyphenyl)octyl]oxy]-2-pyridinyl]methyl]thio]methyl]), Searle SC-53228 (2H-1-Benzopyran-2-propanoic acid, 7-[3-[2-(cyclopropylmethyl) -3-methoxy-4-[(methylamino)carbonyl]phenoxy]propoxy]-3,4-dihydro-8-propyl-, (2S)), Sumitamo SM 15178 (.beta.-Alanine, N-[[6-[(4-acetyl-2ethyl-5-hydroxyphenoxy)methyl]-2- pyridinyl]carbonyl]-N-ethyl), American Home Products WAY 121006 ([1,1'-Biphenyl]-4-acetic acid, 2-fluoro-4'-(2-quinolinylmethoxy)), Bayer Bay-o-8276, calcitriol (9,10-Secocholesta-5,7,10(19)-triene-1,3,25-triol, (1.alpha., 3.beta., 5Z, 7E)), Warner-Lambert CI-987 (2,4-Thiazolidinedione, 5-[[3,5-bis(1,1-dimethylethyl)-4hydroxyphenyl]methylene]), Merck and Co. L-651392 (3H-Phenothiazin-3-one, 4-bromo-2,7-dimethoxy), Lilly LY 210073, Lilly LY 223982 (Benzenepropanoic acid, 5-(3-carboxybenzoyl)-2-[[(5E)-6-(4-methoxyphenyl)-5- hexenyl]oxy]), Lilly LY-233569 (2-Propenamide, N-hydroxy-N-methyl-3-[2-(methylthio)phenyl]), Lilly LY-255283 (Ethanone, 1-[5-ethyl-2-hydroxy-4-[[6-methyl-6-(1Htetrazol-5-yl)heptyl]oxy]phenyl]), Merck and Co. MK-591 (1H-Indole-2-propanoic acid, 1-[(4-chlorophenyl)methyl]-3-[(1,1dimethylethyl)thio]-a,a-dimethyl-5-(2-quinolinylmethoxy)-, sodium salt), Merck and CO. MK-886 (1H-indole-2-propanoic acid, 1-[(4chlorophenyl)methyl]-3-[(1,1-dimethylethyl)thio]-a,a-dimethyl-5-(1-methylethyl)), Ono ONO-LB-448, Purdue Frederick PF-5901 (Benzenemethanol, a-pentyl-3-(2-quinolinylmethoxy)), Rhone-Poulenc Rorer RG 14893 (2-Naphthalenecarboxylic acid, 4-[2-



[methyl(2-phenylethyl) amino] -2-oxoethyl] -8-(phenylmethoxy)),
Rhone-Poulenc Rorer RP 66364, Rhone-Poulenc Rorer RP 69698
(Pyridine, 2-[[5-methyl-5-(1H-tetrazol-5-yl)hexyl]oxy] -4,6diphenyl), Searle SC-41930 (2H-1-Benzopyran-2-carboxylic acid, 7[3-(4-acetyl-3-methoxy-2-propylphenoxy)propoxy] -3,4-dihydro-8propyl), Searle SC-50505, Searle SC-51146, SmithKline Beecham
SK&F-104493 (5H-Pyrrolo[1,2-a]imidazole, 6,7-dihydro-2-(4methoxyphenyl) -3-(4-pyridinyl)), and Teinjin TEI-1338 (Benzoic acid, 2-[[4-[2-[2-(2-naphthalenyl)ethenyl]cyclopropyl]-1oxobutyl]amino]-, methyl ester, [1R-[1.alpha.,2.beta.(E)]]).

### CLAIM 4:

4. (once amended) The combination of Claim 3 wherein the leukotriene B4 receptor antagonist is selected from Bayer Bay-x-1005 ((R)-a-Cyclopentyl-4-(2-quinolinylmethoxy)benzeneacetic acid), Ciba-Geigy CGS-25019C (Benzamide, 4-[[5-[4-(aminoiminomethyl) phenoxy | pentyl | oxy | -3-methoxy-N, N-bis(1methylethyl) -, (2Z) -2-butenedioate), ebselen (1,2-Benzisoselenazol-3(2H)-one, 2-phenyl), Leo Denmark ETH-615 (Benzoic acid, 4-[[[(3-fluorophenyl)methyl][4-(2quinolinylmethoxy)phenyl]amino]methyl]), Lilly LY-293111 (Benzoic acid, 2-[3-[3-[(5-ethyl-4'-fluoro-2-hydroxy[1,1'-biphenyl]-4yl)oxy]propoxy]-2-propylphenoxy]), Ono ONO-4057 (Benzenepropanoic acid, 2-(4-carboxybutoxy)-6-[[(5E)-6-(4-methoxyphenyl)-5hexenyl]oxy]), Terumo TMK-688 (Carbonic acid, 4-[5-[[2-[4-(diphenylmethoxy) -1-piperidinyl]ethyl]amino]-5-oxo-1,3pentadienyl]-2-methoxyphenyl ethyl ester), Boehringer Ingleheim BI-RM-270 (2-Benzoxazolamine, N-[(1S)-2-cyclohexyl-1-(2pyridinyl)ethyl]-5-methyl), Lilly LY 213024 (Benzenepropanoic acid, 5-(3-carboxybenzoyl)-2-(decyloxy)), Lilly LY 264086 (9H-Xanthene-4-propanoic acid, 7-carboxy-3-(decyloxy)-9-oxo), Lilly LY 292728, Ono ONO LB457 (Benzenepropanoic acid, 2-(4carboxybutoxy) -6-[[(5E) -6-(4-methoxyphenyl) -5-hexenyl]oxy]), Pfizer 105696, Perdue Frederick PF 10042 (Pyrrolidine, 1-[5hydroxy-5-[8-(1-hydroxy-2-phenylethyl)-2-dibenzofuranyl]-1oxopentyl]), Rhone-Poulenc Rorer RP 66153 (2-Thiopheneheptanoic



acid, .alpha.,.alpha.-dimethyl-3-(3-phenylpropyl)), SmithKline
Beecham SB-201146 (2-Propenoic acid, 3-[6-[[(3-aminophenyl)sulfinyl]methyl]-3-[[8-(4-methoxyphenyl)octyl]oxy]2-pyridinyl]-, (2E)), SmithKline Beecham SB-201993 (Benzoic acid,
3-[[[6-[(1E)-2-carboxyethenyl]-5-[[8-(4-methoxyphenyl)octyl]oxy]-2-pyridinyl]methyl]thio]methyl]), Searle
SC-53228 (2H-1-Benzopyran-2-propanoic acid, 7-[3-[2-(cyclopropylmethyl)-3-methoxy-4[(methylamino)carbonyl]phenoxylpropoxyl-3,4-dihydro-8-propyl-,
(2S)), Sumitamo SM 15178 (.beta.-Alanine, N-[[6-[(4-acetyl-2-ethyl-5-hydroxyphenoxy)methyl]-2- pyridinyl]carbonyl]-N-ethyl),
and American Home Products WAY 121006 ([1,1'-Biphenyl]-4-acetic acid, 2-fluoro-4'-(2-quinolinylmethoxy)).

#### CLAIM 5:

5. (once amended) The combination of Claim 4 wherein the leukotriene B<sub>4</sub> receptor antagonist is selected from Bayer Bay-x-1005 ((R)-a-Cyclopentyl-4-(2-quinolinylmethoxy)benzeneacetic acid), Ciba-Geigy CGS-25019C (Benzamide, 4-[[5-[4-(aminoiminomethyl)phenoxy]pentyl]oxy]-3-methoxy-N,N-bis(1-methylethyl)-, (2Z)-2-butenedioate), ebselen (1,2-Benzisoselenazol-3(2H)-one, 2-phenyl), Leo Denmark ETH-615 (Benzoic acid, 4-[[(3-fluorophenyl)methyl][4-(2-quinolinylmethoxy)phenyl]amino]methyl]), Lilly LY-293111 (Benzoic acid, 2-[3-[3-[(5-ethyl-4'-fluoro-2-hydroxy[1,1'-biphenyl]-4-yl)oxy]propoxy]-2-propylphenoxy]), Ono ONO-4057 (Benzenepropanoic acid, 2-(4-carboxybutoxy)-6-[[(5E)-6-(4-methoxyphenyl)-5-hexenyl]oxy]), and Terumo TMK-688 (Carbonic acid, 4-[5-[[2-[4-(diphenylmethoxy)-1-piperidinyl]ethyl]amino]-5-oxo-1,3-pentadienyl]-2-methoxyphenyl ethyl ester).

### CLAIM 9:

9. (once amended) A pharmaceutical composition comprising a pharmaceutically-acceptable carrier and a therapeutically-effective amount of a leukotriene B<sub>4</sub> receptor antagonist and a cyclooxygenase-2 inhibitor selected from Taisho NS-398

2 /3 (PHA 4142.2) PATENT

(Methanesulfonamide, N-[2-(cyclohexyloxy)-4-nitrophenyl]),
meloxicam (2H-1,2-Benzothiazine-3-carboxamide, 4-hydroxy-2methyl-N-(5-methyl-2-thiazolyl)-, 1,1-dioxide), [floculide]
flosulide (Methanesulfonamide, N-[6-(2,4-difluorophenoxy)-2,3dihydro-1-oxo-1H-inden-5-yl]), Merck MK-966 (2(5H)-Furanone, 4[4-(methylsulfonyl)phenyl]-3-phenyl), Merck L-752,860 and
compounds of Formula I

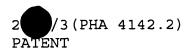
$$R^2$$
  $R^2$   $R^2$   $R^2$ 

wherein A is a substituent selected from partially unsaturated or unsaturated heterocyclyl and partially unsaturated or unsaturated carbocyclic rings;

wherein R<sup>1</sup> is at least one substituent selected from heterocyclyl, cycloalkyl, cycloalkenyl and aryl, wherein R<sup>1</sup> is optionally substituted at a substitutable position with one or more radicals selected from alkyl, haloalkyl, cyano, carboxyl, alkoxycarbonyl, hydroxyl, hydroxyalkyl, haloalkoxy, amino, alkylamino, arylamino, nitro, alkoxyalkyl, alkylsufinyl, halo, alkoxy and alkylthio;

wherein  $R^2$  is methyl or amino; and

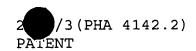
wherein R³ is a radical selected from hydrido, halo, alkyl, alkenyl, alkynyl, oxo, cyano, carboxyl, cyanoalkyl, heterocyclyloxy, alkyloxy, alkylthio, alkylcarbonyl, cycloalkenyl, aralkyl, heterocyclylalkyl, acyl, alkylthioalkyl, hydroxyalkyl, alkoxycarbonyl, arylcarbonyl, aralkylcarbonyl, aralkenyl, alkoxyalkyl, arylthioalkyl, aryloxyalkyl, aralkylthioalkyl, aralkoxyalkyl, alkoxyaralkoxyalkyl, alkoxycarbonylalkyl, aminocarbonyl, aminocarbonylalkyl, alkylaminocarbonyl, N-alkyl-N-arylaminocarbonyl, N-alkyl-N-aralkylamino, N-arylamino, N-aralkylamino, N-alkyl-N-aralkylamino, N-arylamino, aminoalkyl, alkylaminoalkyl, N-arylaminoalkyl, N-aralkylaminoalkyl, N-aralkylaminoalkyl,



N-alkyl-N-arylaminoalkyl, aryloxy, aralkoxy, arylthio, aralkylthio, alkylsulfinyl, alkylsulfonyl, aminosulfonyl, alkylaminosulfonyl, N-arylaminosulfonyl, arylsulfonyl, N-alkyl-N-arylaminosulfonyl;

or a pharmaceutically-acceptable salt thereof.

Please cancel claims 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22 and 23.



charge Deposit Account No. 19-1345.

Respectfully submitted,

Kathryn J. Doty, Reg. No. 40 593 SENNIGER, POWERS, LEAVITT & ROEDEL One Metropolitan Square, 16th Floor

St. Louis, Missouri 63102

(314) 231-5400

Express Mail No.: EL 801513198 US